

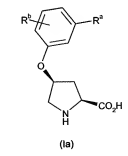
Listing of Claims

The following listing of Claims will replace all prior versions and listings of Claims in the Application.

Claims

1-10 (Cancelled)

11. (Currently Amended) A pharmaceutical composition comprising a compound of formula (1a) according to claim 14 :

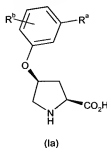


wherein R<sup>a</sup> is selected from halogen, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, cyano, nitro, amino, hydroxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkenyl, C<sub>1</sub>-C<sub>6</sub> alkynyl, hydroxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkoxyC<sub>1</sub>-C<sub>6</sub> alkyl, perfluoro C<sub>1</sub>-C<sub>6</sub> alkyl, perfluoroC<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylamino, di- C<sub>1</sub>-C<sub>6</sub> alkylamino, aminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acyl, C<sub>1</sub>-C<sub>6</sub> acyloxy, C<sub>1</sub>-C<sub>6</sub> acyloxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acylamino, C<sub>1</sub>-C<sub>6</sub> alkylthiocarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkoxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonylamino, aminosulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, 3-8 membered cycloalkyl, 4-8 membered heterocycloalkyl, phenyl and monocyclic heteroaryl;

R<sup>b</sup> is selected from hydrogen, halogen, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy cyano, nitro, amino, hydroxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkenyl, C<sub>1</sub>-C<sub>6</sub> alkynyl,

C<sub>1</sub>-C<sub>6</sub> alkoxy, hydroxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkoxyC<sub>1</sub>-C<sub>6</sub> alkyl, perfluoro C<sub>1</sub>-C<sub>6</sub> alkyl, perfluoroC<sub>1</sub>-C<sub>6</sub> alkoxy,  
C<sub>1</sub>-C<sub>6</sub> alkylamino, di- C<sub>1</sub>-C<sub>6</sub> alkylamino, aminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl,  
C<sub>1</sub>-C<sub>6</sub> acyl, C<sub>1</sub>-C<sub>6</sub> acyloxy, C<sub>1</sub>-C<sub>6</sub> acyloxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acylamino,  
C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkylthiocarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkoxy carbonyl,  
C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonylamino,  
aminosulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl,  
3-8 membered cycloalkyl, 4-8 membered heterocycloalkyl, phenyl and  
monocyclic heteroaryl;  
 or a pharmaceutically acceptable salt thereof, and one or more pharmaceutically acceptable excipients, diluents or carriers.

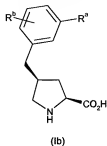
12. (Previously Presented) A combination comprising a compound of formula (Ia) according to claim 14, or a pharmaceutically acceptable salt thereof, and at least one other therapeutically active agent.
13. (Currently Amended) A combination according to claim 12, wherein the other therapeutically active agent is a PDEV inhibitor selected from sildenafil, vardenafil, tadalafil, 1-[6-ethoxy-5-[3-ethyl-6,7-dihydro-2-(2-methoxyethyl)-7-oxo-2H-pyrazolo[4,3- d]pyrimidin-5-yl]-3-pyridylsulfonyl]-4-ethylpiperazine, 5-(5-acetyl-2-butoxy-3-pyridinyl)-3-ethyl-2-(1-ethyl-3-azetidinyl)-2,6-dihydro-7H-pyrazolo[4,3-d]pyrimidin-7-one and 5-[2-ethoxy-5-(4-ethylpiperazin-1-ylsulphonyl)pyridin-3-yl]-3-ethyl-2-[2-methoxyethyl]-2,6-dihydro-7H-pyrazolo[4,3-d]pyrimidin-7-one.
14. (Previously Presented) A compound of formula (Ia):



wherein R<sup>a</sup> is selected from halogen, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, cyano, nitro, amino, hydroxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkenyl, C<sub>1</sub>-C<sub>6</sub> alkynyl, hydroxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkoxyC<sub>1</sub>-C<sub>6</sub> alkyl, perfluoro C<sub>1</sub>-C<sub>6</sub> alkyl, perfluoroC<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylamino, di- C<sub>1</sub>-C<sub>6</sub> alkylamino, aminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acyl, C<sub>1</sub>-C<sub>6</sub> acyloxy, C<sub>1</sub>-C<sub>6</sub> acyloxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acylamino, C<sub>1</sub>-C<sub>6</sub> alkylthiocarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkoxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonylamino, aminosulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, 3-8 membered cycloalkyl, 4-8 membered heterocycloalkyl, phenyl and monocyclic heteroaryl;

R<sup>b</sup> is selected from hydrogen, halogen, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy cyano, nitro, amino, hydroxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkenyl, C<sub>1</sub>-C<sub>6</sub> alkynyl, C<sub>1</sub>-C<sub>6</sub> alkoxy, hydroxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkoxyC<sub>1</sub>-C<sub>6</sub> alkyl, perfluoro C<sub>1</sub>-C<sub>6</sub> alkyl, perfluoroC<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylamino, di- C<sub>1</sub>-C<sub>6</sub> alkylamino, aminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acyl, C<sub>1</sub>-C<sub>6</sub> acyloxy, C<sub>1</sub>-C<sub>6</sub> acyloxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acylamino, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkylthiocarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkoxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonylamino, aminosulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, 3-8 membered cycloalkyl, 4-8 membered heterocycloalkyl, phenyl and monocyclic heteroaryl; or a pharmaceutically acceptable salt thereof.

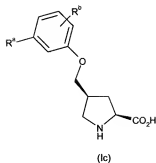
15. (Withdrawn) A compound of formula (Ib):



wherein R<sup>a</sup> is selected from halogen, hydroxy, cyano, nitro, amino, hydroxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkenyl, C<sub>1</sub>-C<sub>6</sub> alkynyl, hydroxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkoxyC<sub>1</sub>-C<sub>6</sub> alkyl, perfluoroC<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylamino, di-C<sub>1</sub>-C<sub>6</sub> alkylamino, aminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub>acyl, C<sub>1</sub>-C<sub>6</sub>acyloxy, C<sub>1</sub>-C<sub>6</sub>acyloxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acylamino, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkylthiocarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylthioxo, C<sub>1</sub>-C<sub>6</sub> alkoxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonylamino, aminosulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, 3-8 membered cycloalkyl, 4-8 membered heterocycloalkyl, phenyl and monocyclic heteroaryl;

R<sup>b</sup> is selected from hydrogen, halogen, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy cyano, nitro, amino, hydroxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkenyl, C<sub>1</sub>-C<sub>6</sub> alkynyl, C<sub>1</sub>-C<sub>6</sub> alkoxy, hydroxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkoxyC<sub>1</sub>-C<sub>6</sub> alkyl, perfluoro C<sub>1</sub>-C<sub>6</sub> alkyl, perfluoroC<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylamino, di-C<sub>1</sub>-C<sub>6</sub> alkylamino, aminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub>acyl, C<sub>1</sub>-C<sub>6</sub>acyloxy, C<sub>1</sub>-C<sub>6</sub>acyloxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acylamino, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkylthiocarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylthioxo, C<sub>1</sub>-C<sub>6</sub> alkoxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonylamino, aminosulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, 3-8 membered cycloalkyl, 4-8 membered heterocycloalkyl, phenyl and monocyclic heteroaryl; or a pharmaceutically acceptable salt thereof.

16. (Withdrawn) A compound of formula (Ic):



wherein  $R^a$  and  $R^b$  are independently selected from hydrogen, halogen, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, cyano, nitro, amino, hydroxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkenyl, C<sub>1</sub>-C<sub>6</sub> alkynyl, C<sub>1</sub>-C<sub>6</sub> alkoxy, hydroxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkoxyC<sub>1</sub>-C<sub>6</sub> alkyl, perfluoro C<sub>1</sub>-C<sub>6</sub> alkyl, perfluoroC<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylamino, di- C<sub>1</sub>-C<sub>6</sub> alkylamino, aminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminoC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acyl, C<sub>1</sub>-C<sub>6</sub>acyloxy, C<sub>1</sub>-C<sub>6</sub>acyloxyC<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> acylamino, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkylthiocarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylthioxo, C<sub>1</sub>-C<sub>6</sub> alkoxycarbonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonylamino, aminosulfonyl, C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, di-C<sub>1</sub>-C<sub>6</sub> alkylaminosulfonyl, 3-8 membered cycloalkyl, 4-8 membered heterocycloalkyl, phenyl and monocyclic heteroaryl; or a pharmaceutically acceptable salt thereof.

17. (Withdrawn, Currently Amended) A compound of formula (1a) according to claim 4 14 which is:

(2S, 4S)-4-(3-Chloro-phenoxy)-pyrrolidine-2-carboxylic acid;

Or a pharmaceutically acceptable salt thereof.

18. (Withdrawn) A compound of formula (1b) according to claim 15 which is selected from the group consisting of:

(2S,4S)-4-(3-Fluoro-benzyl)-pyrrolidine-2-carboxylic acid;

(2S,4S)-4-(2,3-Difluoro-benzyl)-pyrrolidine-2-carboxylic acid; and

(2S,4S)-4-(2,5-Difluoro-benzyl)-pyrrolidine-2-carboxylic acid; or a pharmaceutically acceptable salt thereof.

19. (Withdrawn) A compound of formula (1c) according to claim 16 which is selected from the group consisting of:

(2S,4S)-4-(3-Fluoro-phenoxy-methyl)-pyrrolidine-2-carboxylic acid;  
(2S,4S)-4-(3,6-Difluoro-phenoxy-methyl)-pyrrolidine-2-carboxylic acid;  
(2S,4S)-4-(2,3-Difluoro-phenoxy-methyl)-pyrrolidine-2-carboxylic acid; and  
(2S,4S)-4-(3-Methoxy- phenoxy-methyl)-pyrrolidine-2-carboxylic acid; or a pharmaceutically acceptable salt thereof.

20. (Withdrawn) A pharmaceutical composition comprising a compound of formula (1b) according to claim 15, or a pharmaceutically acceptable salt thereof, and one or more pharmaceutically acceptable excipients, diluents or carriers.

21. (Withdrawn) A pharmaceutical composition comprising a compound of formula (1b) according to claim 15, or a pharmaceutically acceptable salt thereof, and at least one other therapeutically active agent.

22. (Withdrawn) A combination according to claim 21, wherein the other therapeutically active agent is a PDEV inhibitor.

23. (Withdrawn) A pharmaceutical composition comprising a compound of formula (1c) according to claim 16, or a pharmaceutically acceptable salt thereof, and one or more pharmaceutically acceptable excipients, diluents or carriers.

24. (Withdrawn) A pharmaceutical composition comprising a compound of formula (1c) according to claim 16, or a pharmaceutically acceptable salt thereof, and at least one other therapeutically active agent.

25. (Withdrawn) A combination according to claim 24, wherein the other therapeutically active agent is a PDEV inhibitor.

26. (Withdrawn) The compound (2S, 4S)-4-(3-Chloro-phenoxy)-pyrrolidine-2-carboxylic acid or a pharmaceutically acceptable salt thereof.